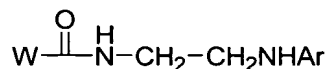


WHAT IS CLAIMED IS:

1. A compound of Formula I:



(I)

or a pharmaceutically acceptable salt or prodrug thereof,
wherein:

W is a member selected from the group consisting of

$\text{R}^1-\text{X}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^2-$,

$\text{R}^4-\text{Y}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^3-$,

$\text{R}^6-(\text{C}=\text{O})-\text{NH}-\text{CHR}^5-$,

$\text{R}^7-\text{NH}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^8-$,

$\text{R}^{10}-\text{Z}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^9-$, and

$\text{R}^{11}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^{12}-$;

R^1 is a member selected from the group consisting of phenyl substituted with 0-2 R^{1a} ,
pyridyl substituted with 0-2 R^{1a} , and pyridinium N-oxide substituted with 0-2
 R^{1a} ;

each R^{1a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

X is a member selected from the group consisting of furanylene substituted with 0-1
 R^x , thienylene substituted with 0-1 R^x , pyrazolyene substituted with 0-1 R^x ,
thiazolyene substituted with 0-1 R^x , and oxazolyene substituted with 0-1 R^x ;

R^x is a member selected from the group consisting of F, Cl, CH_3 and CF_3 ;

R^2 is a member selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
and $(\text{CH}_2)_n\text{R}^{2b}$;

each R^{2a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

R^{2b} is independently a member selected from the group consisting of phenyl
substituted with 0-2 R^{2a} ; cyclopentyl, cyclohexyl and tetrahydropyranyl;

n is the integer 1 or 2;

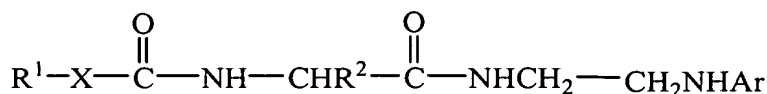
R^3 is $(\text{CH}_2)_m\text{R}^{3b}$;

R^{3b} is selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
cyclopentyl and cyclohexyl;
m is the integer 1 or 2;
 R^4 is a member selected from the group consisting of phenyl substituted with 0-3 R^{4a} ,
thienyl, tetrazolyl, cyclopentenyl and indolyl;
each R^{4a} is a member selected from the group consisting of phenyl, OH, C_1 - C_4 alkyl,
 C_1 - C_4 alkoxy, CF_3 , OCF_3 , F, Cl, $CH_3S(=O)_2$ -, morpholinyl, pyrrolidinyl,
piperidinyl and 4-acetylpiperazinyl;
Y is a member selected from the group consisting of $-CR^{17}R^{18}$, $-NH-CH_2-$ and
 $-O-CH_2-$;
 R^5 is a member selected from the group consisting of phenyl substituted with 0-2 R^{5a} ,
thiophene, naphthyl, and CH_2R^{5b} , CH_2CH_2 (cyclohexyl),
 $CH_2CH_2CH_2$ (cyclohexyl), CH_2CH_2Ph , $CH(CH_3)R^{5c}$, $CH_2CH=CHPh$, -
 CH_2OCH_2Ph , $-CH(CH_3)OCH_2Ph$;
each R^{5a} is independently a member selected from the group consisting of F, Cl, NO_2 ,
 OCH_3 , OCH_2Ph , OPh , CH_3 , OCF_3 and CF_3 ;
 R^{5b} is independently a member selected from the group consisting of phenyl
substituted with 0-2 R^{5c} ; cyclopentyl, cyclohexyl, naphthyl, indolyl and
pyridyl;
 R^{5c} is independently a member selected from the group consisting of OH, Cl, F, Br, I,
CN, NO_2 , CH_3 , OCH_3 , tBu , $O-tBu$, $-NHC(=O)CH_3$, CF_3 , OCF_3 ; phenyl
substituted with 0-2 R^{5d} ; phenoxy substituted with 0-2 R^{5d} ; benzyloxy
substituted with 0-2 R^{5d} ; pyridyl substituted with 0-2 R^{5d} ; pyrimidinyl
substituted with 0-2 R^{5d} ; thienyl substituted with 0-2 R^{5d} ;
 R^{5d} is independently a member selected from the group consisting of CH_3 , Cl, F,
 OCH_3 , CF_3 , OCF_3 , $N(CH_3)_2$, acetyl, OH, CH_2OH , NH_2 , CN and NO_2 ;
 R^{5e} is phenyl substituted with 0-2 R^{5a} ;
 R^6 is a member selected from the group consisting of phenyl substituted with 0-3 R^{6a} ,
furanyl substituted with 0-2 R^{6b} , thienyl substituted with 0-2 R^{6b} , oxazolyl
substituted with 0-2 R^{6b} , thiazolyl substituted with 0-2 R^{6b} , pyridyl,
pyridazinyl and cyclopropyl;
each R^{6a} is independently a member selected from the group consisting of Cl, F, Br,
 OCF_3 , CF_3 , C_1 - C_4 alkyl, C_1 - C_4 alkoxy, $-S(=O)_2CH_3$, CN, $-N(CH_3)_2$, OCF_2H , -

63 CH₂-benzimidazole, -NH-S(=O)₂CH₃, -NR¹³R¹⁴, OR¹⁴, CH₂-morpholine,
 64 CH₂NH₂, OCH₂Ph, and OPh;
 65 alternatively, two R^{6a} substituents on adjacent atoms may be combined to form a 5 to
 66 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 67 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 68 each R^{6b} is independently a member selected from the group consisting of NH₂, F, Cl,
 69 Br, -S(=O)₂R¹⁵, CH₃, and CF₃;
 70 R⁷ is a member selected from the group consisting of (CH₂)_p R^{7a}, and naphthyl
 71 substituted with 0-2 R^{7b};
 72 p is the integer 0, 1, or 2;
 73 R^{7a} is phenyl substituted with 0-2 R^{7b};
 74 R^{7b} is a member selected from the group consisting of F, Cl, CF₃, C₁-C₄ alkyl, C₁-C₄
 75 alkoxy, OCF₃, phenoxy and acetyl;
 76 alternatively, two R^{7b} substituents on adjacent atoms may be combined to form a 5 to
 77 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 78 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 79 R⁸ is -CH₂-R^{3b};
 80 R⁹ is (CH₂)_qR^{9a};
 81 R^{9a} is a member selected from the group consisting of cyclopentyl, phenyl and
 82 cyclohexyl;
 83 q is the integer 1 or 2;
 84 R¹⁰ is a member selected from the group consisting of phenyl substituted with 0-2
 85 R^{10a}, 5 membered heteroaryl containing 1 to 4 heteroatoms each independently
 86 a member selected from the group consisting of N, O and S, wherein said
 87 heteroaryl is substituted with 0-2 R^{10a}, 6 membered heteroaryl containing 1 to
 88 2 N, wherein said heteroaryl is substituted with 0-2 R^{10a}, morpholinyl
 89 substituted with 0-2 R^{10a}, piperazinyl substituted with 0-2 R^{10a} and piperidinyl
 90 substituted with 0-2 R^{10a};
 91 each R^{10a} is independently a member selected from the group consisting of Cl, F, C₁-
 92 C₄ alkyl, C₁-C₄ alkoxy, OCF₃, and CF₃;
 93 alternatively, two R^{10a} substituents on adjacent atoms may be combined to form a 5 to
 94 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 95 heterocyclic fused radical comprises 1 or 2 heteroatom(s);
 96 Z is phenylene;

R^{11} is a member selected from the group consisting of indolyl substituted with 0-2 R^{11a} , benzofuranyl substituted with 0-2 R^{11a} , benzothienyl substituted with 0-2 R^{11a} , benzothiazole substituted with 0-2 R^{11a} , benzisoxazolyl substituted with 0-2 R^{11a} , benzoxazolyl substituted with 0-2 R^{11a} , and pyrazolo[1,5-a]pyrimidinyl substituted with 0-2 R^{11a} , piperidinyl N-substituted with 0-1 R^{11b} , morpholinyl N-substituted with 0-1 R^{11b} ; and 2-oxo-pyrrolidinyl with 0-1 R^{11b} ;
each R^{11a} is independently a member selected from the group consisting of Cl, F, NH_2 , CH_3 , OCH_3 , $-C(=O)OCH_3$, OCF_3 , and CF_3 ;
each R^{11b} is independently a member selected from the group consisting of pyrimidinyl substituted with 0-2 R^{11c} ; benzyl, acetyl, CH_2 -furanyl, and CH_2 -thienyl;
each R^{11c} is independently a member selected from the group consisting of Br and CH_3 ;
 R^{12} is $(CH_2)_sR^{12a}$;
 R^{12a} is a member selected from the group consisting of cyclopentyl and cyclohexyl;
s is the integer 1 or 2;
 R^{13} is a member selected from the group consisting of H and C_1 - C_4 alkyl;
 R^{14} is pyrimidinyl substituted with 0-2 R^{16} ;
 R^{15} is a member selected from the group consisting of C_1 - C_4 alkyl, morpholinyl, pyrrolidinyl and piperidinyl;
 R^{16} is a member selected from the group consisting of CH_3 and OCH_3 ;
each of R^{17} and R^{18} is independently a member of H, OH, F, phenyl and C_1 - C_3 alkyl; alternatively, R^{17} and R^{18} may be taken together to form a C_3 - C_6 cycloalkyl;
Ar is a phenyl substituted with 0-2 R^{19} ; and
each R^{19} is independently a member selected from the group consisting of F, Cl, $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 .

2. The compound of claim 1, wherein said compound has the formula:

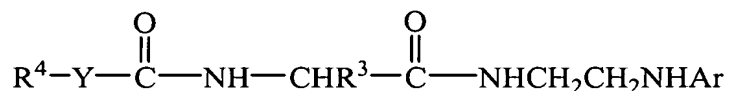


Ia

wherein:

R^1 is a member selected from the group consisting of phenyl substituted with 0-2 R^{1a} ,
 pyridyl substituted with 0-2 R^{1a} , and pyridinium N-oxide substituted with 0-2
 R^{1a} ;
 each R^{1a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;
 X is a member selected from the group consisting of furanylene substituted with 0-1
 R^x , thienylene substituted with 0-1 R^x , pyrazolyne substituted with 0-1 R^x ,
 thiazolyne substituted with 0-1 R^x , and oxazolyne substituted with 0-1 R^x ;
 R^x is a member selected from the group consisting of F, Cl, CH_3 and CF_3 ;
 R^2 is a member selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
 and $(CH_2)_nR^{2b}$;
 each R^{2a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;
 R^{2b} is independently a member selected from the group consisting of phenyl
 substituted with 0-2 R^{2a} , cyclopentyl, cyclohexyl and tetrahydropyranyl;
 n is the integer 1 or 2;
 Ar is a phenyl substituted with 0-2 R^{19} ; and
 each R^{19} is independently a member selected from the group consisting of F, Cl,
 $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 .

3. The compound of claim 1, wherein said compound has the formula:



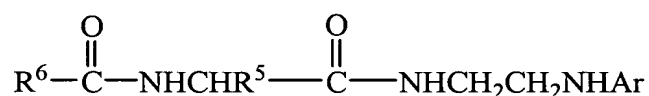
Ib

wherein:

R^4 is a member selected from the group consisting of phenyl substituted with 0-3 R^{4a} ,
 thienyl, tetrazolyl, cyclopentenyl and indolyl;
 each R^{4a} is a member selected from the group consisting of phenyl, OH, C_1 - C_4 alkyl,
 C_1 - C_4 alkoxy, CF_3 , OCF_3 , F, Cl, $CH_3S(=O)_2$ -, morpholinyl, pyrrolidinyl,
 piperidinyl and 4-acetylpiperazinyl;
 Y is a member selected from the group consisting of $-CR^{17}R^{18}$, $-NH-CH_2-$ and
 $-O-CH_2-$;
 R^3 is $(CH_2)_mR^{3b}$;

R^{3b} is selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
 cyclopentyl and cyclohexyl;
 each R^{2a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;
 m is the integer 1 or 2;
 each of R^{17} and R^{18} is independently a member of H, OH, F, phenyl and C_1 - C_3 alkyl;
 alternatively, R^{17} and R^{18} may be taken together to form a C_3 - C_6 cycloalkyl;
 Ar is a phenyl substituted with 0-2 R^{19} ; and
 each R^{19} is independently a member selected from the group consisting of F, Cl,
 $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 .

4. The compound of claim 1, wherein said compound has the formula:



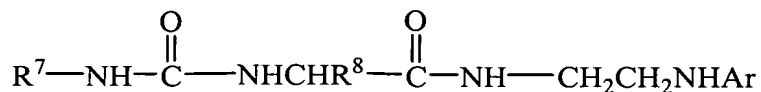
Ic

wherein:

R^5 is a member selected from the group consisting of phenyl substituted with 0-2 R^{5a} ,
 thiophene, naphthyl, and CH_2R^{5b} , CH_2CH_2 (cyclohexyl),
 $CH_2CH_2CH_2$ (cyclohexyl), CH_2CH_2Ph , $CH(CH_3)R^{5c}$, $CH_2CH=CHPh$, -
 CH_2OCH_2Ph , and $-CH(CH_3)OCH_2Ph$;
 each R^{5a} is independently a member selected from the group consisting of F, Cl, NO_2 ,
 OCH_3 , OCH_2Ph , OPh , CH_3 , OCF_3 and CF_3 ;
 R^{5b} is independently a member selected from the group consisting of phenyl
 substituted with 0-2 R^{5c} ; cyclopentyl, cyclohexyl, naphthyl, indolyl and
 pyridyl;
 R^{5c} is independently a member selected from the group consisting of OH, Cl, F, Br, I,
 CN, NO_2 , CH_3 , OCH_3 , tBu , $O-tBu$, $-NHC(=O)CH_3$, CF_3 , OCF_3 , phenyl
 substituted with 0-2 R^{5d} , phenoxy substituted with 0-2 R^{5d} , benzyloxy
 substituted with 0-2 R^{5d} , pyridyl substituted with 0-2 R^{5d} , pyrimidinyl
 substituted with 0-2 R^{5d} , and thienyl substituted with 0-2 R^{5d} ;
 R^{5d} is independently a member selected from the group consisting of CH_3 , Cl, F,
 OCH_3 , CF_3 , OCF_3 , $N(CH_3)_2$, acetyl, OH, CH_2OH , NH_2 , CN and NO_2 ;
 R^{5e} is phenyl substituted with 0-2 R^{5a} ;

R^6 is a member selected from the group consisting of phenyl substituted with 0-3 R^{6a} ,
 furanyl substituted with 0-2 R^{6b} ; thienyl substituted with 0-2 R^{6b} ; oxazolyl
 substituted with 0-2 R^{6b} ; thiazolyl substituted with 0-2 R^{6b} ; pyridyl,
 pyridazinyl and cyclopropyl;
 each R^{6a} is independently a member selected from the group consisting of Cl, F, Br,
 OCF_3 , CF_3 , C_1 - C_4 alkyl, C_1 - C_4 alkoxy, $-S(=O)_2CH_3$, CN, $-N(CH_3)_2$, OCF_2H , $-$
 CH_2 -benzimidazole, $-NH-S(=O)_2CH_3$, $-NR^{13}R^{14}$, OR^{14} , CH_2 -morpholine,
 CH_2NH_2 , OCH_2Ph , and OPh ;
 alternatively, two R^{6a} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 each R^{6b} is independently a member selected from the group consisting of NH_2 , F, Cl,
 Br, $-S(=O)_2R^{15}$, CH_3 , and CF_3 ;
 R^{13} is a member selected from the group consisting of H and C_1 - C_4 alkyl;
 R^{14} is pyrimidinyl substituted with 0-2 R^{16} ;
 R^{15} is a member selected from the group consisting of C_1 - C_4 alkyl, morpholinyl,
 pyrrolidinyl and piperidinyl;
 R^{16} is a member selected from the group consisting of CH_3 and OCH_3 ;
 Ar is a phenyl substituted with 0-2 R^{19} ; and
 each R^{19} is independently a member selected from the group consisting of F, Cl,
 $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 .

5. The compound of claim 1, wherein said compound has the formula:



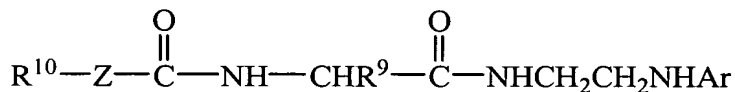
Id

wherein:

R^7 is a member selected from the group consisting of $(CH_2)_p R^{7a}$; and naphthyl
 substituted with 0-2 R^{7b} ;
 p is the integer 0, 1, or 2;
 R^{7a} is a member selected from the group consisting of phenyl substituted with 0-2 R^{7b} ;
 R^{7b} is a member selected from the group consisting of F, Cl, CF_3 , C_1 - C_4 alkyl, C_1 - C_4
 alkoxy, OCF_3 , phenoxy and acetyl;

alternatively, two R^{7b} substituents on adjacent atoms may be combined to form a 5 to 6 membered heterocyclic fused radical, wherein said 5 to 6 membered heterocyclic fused radical has 1 or 2 oxygen atom(s);
R⁸ is -CH₂-R^{3b};
R^{3b} is selected from the group consisting of phenyl substituted with 0-2 R^{2a}, cyclopentyl and cyclohexyl;
each R^{2a} is independently a member selected from the group consisting of Cl, F, OCF₃, OCH₃, CH₃ and CF₃;
Ar is a phenyl substituted with 0-2 R¹⁹; and
each R¹⁹ is independently a member selected from the group consisting of F, Cl, COOH, C₁-C₄ alkoxy, OCHF₂ and OCF₃.

6. The compound of claim 1, wherein said compound has the formula:



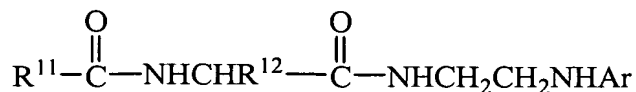
Ie

wherein:

R¹⁰ is a member selected from the group consisting of phenyl substituted with 0-2 R^{10a}, 5 membered heteroaryl containing 1 to 4 heteroatoms each independently a member selected from the group consisting of N, O and S, wherein said heteroaryl is substituted with 0-2 R^{10a}, 6 membered heteroaryl containing 1 to 2 N, wherein said heteroaryl is substituted with 0-2 R^{10a}, morpholinyl substituted with 0-2 R^{10a}, piperazinyl substituted with 0-2 R^{10a} and piperidinyl substituted with 0-2 R^{10a};
each R^{10a} is independently a member selected from the group consisting of Cl, F, C₁-C₄ alkyl, C₁-C₄ alkoxy, OCF₃, and CF₃;
alternatively, two R^{10a} substituents on adjacent atoms may be combined to form a 5 to 6 membered heterocyclic fused radical, wherein said 5 to 6 membered heterocyclic fused radical comprises 1 or 2 heteroatom(s);
Z is phenylene;
R⁹ is (CH₂)_qR^{9a};
R^{9a} is a member selected from the group consisting of cyclopentyl, phenyl and cyclohexyl;

q is the integer 1 or 2;
 Ar is a phenyl substituted with 0-2 R¹⁹; and
 each R¹⁹ is independently a member selected from the group consisting of F, Cl,
 COOH, C₁-C₄ alkoxy, OCHF₂ and OCF₃.

7. The compound of claim 1, wherein said compound has the formula:



If

wherein:

R¹¹ is a member selected from the group consisting of indolyl substituted with 0-2 R^{11a}; benzofuranyl substituted with 0-2 R^{11a}; benzothienyl substituted with 0-2 R^{11a}; benzothiazole substituted with 0-2 R^{11a}; benzisoxazolyl substituted with 0-2 R^{11a}; benzoxazolyl substituted with 0-2 R^{11a}; and pyrazolo[1,5-a]pyrimidinyl substituted with 0-2 R^{11a}; piperidinyl N-substituted with 0-1 R^{11b}; morpholinyl N-substituted with 0-1 R^{11b}; and 2-oxo-pyrrolidinyl with 0-1 R^{11b};

each R^{11a} is independently a member selected from the group consisting of Cl, F, NH₂, CH₃, OCH₃, -C(=O)OCH₃, OCF₃, and CF₃;

each R^{11b} is independently a member selected from the group consisting of pyrimidinyl substituted with 0-2 R^{11c}; benzyl, acetyl, CH₂-furanyl, and CH₂-thienyl;

each R^{11c} is independently a member selected from the group consisting of Br and CH₃;

R¹² is (CH₂)_sR^{12a};

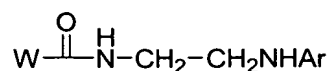
R^{12a} is a member selected from the group consisting of cyclopentyl and cyclohexyl;
 s is the integer 1 or 2;

Ar is a phenyl substituted with 0-2 R¹⁹; and

each R¹⁹ is independently a member selected from the group consisting of F, Cl, COOH, C₁-C₄ alkoxy, OCHF₂ and OCF₃.

8. The compound of claim 1, wherein said compound is a member selected from the compounds of Table I.

9. A pharmaceutical composition, said composition comprising a compound of Formula I:



(I)

or a pharmaceutically acceptable salt or prodrug thereof,
wherein:

W is a member selected from the group consisting of

$\text{R}^1-\text{X}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^2-$,

$\text{R}^4-\text{Y}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^3-$,

$\text{R}^6-(\text{C}=\text{O})-\text{NH}-\text{CHR}^5-$,

$\text{R}^7-\text{NH}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^8-$,

$\text{R}^{10}-\text{Z}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^9-$, and

$\text{R}^{11}-(\text{C}=\text{O})-\text{NH}-\text{CHR}^{12}-$;

R^1 is a member selected from the group consisting of phenyl substituted with 0-2 R^{1a} ,
pyridyl substituted with 0-2 R^{1a} , and pyridinium N-oxide substituted with 0-2
 R^{1a} ;

each R^{1a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

X is a member selected from the group consisting of furanylene substituted with 0-1
 R^x , thienylene substituted with 0-1 R^x , pyrazolyene substituted with 0-1 R^x ,
thiazolyene substituted with 0-1 R^x , and oxazolyene substituted with 0-1 R^x ;

R^x is a member selected from the group consisting of F, Cl, CH_3 and CF_3 ;

R^2 is a member selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
and $(\text{CH}_2)_n\text{R}^{2b}$;

each R^{2a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

R^{2b} is independently a member selected from the group consisting of phenyl
substituted with 0-2 R^{2a} ; cyclopentyl, cyclohexyl and tetrahydropyranyl;

n is the integer 1 or 2;

R^3 is $(\text{CH}_2)_m\text{R}^{3b}$;

R^{3b} is selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
cyclopentyl and cyclohexyl;

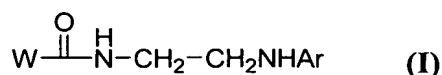
m is the integer 1 or 2;
 R^4 is a member selected from the group consisting of phenyl substituted with 0-3 R^{4a} ,
 thienyl, tetrazolyl, cyclopentenyl and indolyl;
 each R^{4a} is a member selected from the group consisting of phenyl, OH, C₁-C₄ alkyl,
 C₁-C₄ alkoxy, CF₃, OCF₃, F, Cl, CH₃S(=O)₂-, morpholinyl, pyrrolidinyl,
 piperidinyl and 4-acetylpiperazinyl;
 Y is a member selected from the group consisting of $-CR^{17}R^{18}$, $-NH-CH_2-$ and $-O-CH_2-$;
 R^5 is a member selected from the group consisting of phenyl substituted with 0-2 R^{5a} ,
 thiophene, naphthyl, and CH_2R^{5b} , CH_2CH_2 (cyclohexyl),
 $CH_2CH_2CH_2$ (cyclohexyl), CH_2CH_2Ph , $CH(CH_3)R^{5c}$, $CH_2CH=CHPh$, -
 CH_2OCH_2Ph , $-CH(CH_3)OCH_2Ph$;
 each R^{5a} is independently a member selected from the group consisting of F, Cl, NO₂,
 OCH₃, OCH₂Ph, OPh, CH₃, OCF₃ and CF₃;
 R^{5b} is independently a member selected from the group consisting of phenyl
 substituted with 0-2 R^{5c} ; cyclopentyl, cyclohexyl, naphthyl, indolyl and
 pyridyl;
 R^{5c} is independently a member selected from the group consisting of OH, Cl, F, Br, I,
 CN, NO₂, CH₃, OCH₃, ^tBu, O-^tBu, $-NHC(=O)CH_3$, CF₃, OCF₃; phenyl
 substituted with 0-2 R^{5d} ; phenoxy substituted with 0-2 R^{5d} ; benzyloxy
 substituted with 0-2 R^{5d} ; pyridyl substituted with 0-2 R^{5d} ; pyrimidinyl
 substituted with 0-2 R^{5d} ; thienyl substituted with 0-2 R^{5d} ;
 R^{5d} is independently a member selected from the group consisting of CH₃, Cl, F,
 OCH₃, CF₃, OCF₃, N(CH₃)₂, acetyl, OH, CH₂OH, NH₂, CN and NO₂;
 R^{5e} is phenyl substituted with 0-2 R^{5a} ;
 R^6 is a member selected from the group consisting of phenyl substituted with 0-3 R^{6a} ,
 furanyl substituted with 0-2 R^{6b} , thienyl substituted with 0-2 R^{6b} , oxazolyl
 substituted with 0-2 R^{6b} , thiazolyl substituted with 0-2 R^{6b} , pyridyl,
 pyridazinyl and cyclopropyl;
 each R^{6a} is independently a member selected from the group consisting of Cl, F, Br,
 OCF₃, CF₃, C₁-C₄ alkyl, C₁-C₄ alkoxy, $-S(=O)_2CH_3$, CN, $-N(CH_3)_2$, OCF₂H, -
 CH_2 -benzimidazole, $-NH-S(=O)_2CH_3$, $-NR^{13}R^{14}$, OR^{14} , CH_2 -morpholine,
 CH_2NH_2 , OCH₂Ph, and OPh;

alternatively, two R^{6a} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 each R^{6b} is independently a member selected from the group consisting of NH₂, F, Cl,
 Br, -S(=O)₂R¹⁵, CH₃, and CF₃;
 R⁷ is a member selected from the group consisting of (CH₂)_p R^{7a}, and naphthyl
 substituted with 0-2 R^{7b};
 p is the integer 0, 1, or 2;
 R^{7a} is phenyl substituted with 0-2 R^{7b};
 R^{7b} is a member selected from the group consisting of F, Cl, CF₃, C₁-C₄ alkyl, C₁-C₄
 alkoxy, OCF₃, phenoxy and acetyl;
 alternatively, two R^{7b} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 R⁸ is -CH₂-R^{3b};
 R⁹ is (CH₂)_qR^{9a};
 R^{9a} is a member selected from the group consisting of cyclopentyl, phenyl and
 cyclohexyl;
 q is the integer 1 or 2;
 R¹⁰ is a member selected from the group consisting of phenyl substituted with 0-2
 R^{10a}, 5 membered heteroaryl containing 1 to 4 heteroatoms each independently
 a member selected from the group consisting of N, O and S, wherein said
 heteroaryl is substituted with 0-2 R^{10a}, 6 membered heteroaryl containing 1 to
 2 N, wherein said heteroaryl is substituted with 0-2 R^{10a}, morpholinyl
 substituted with 0-2 R^{10a}, piperazinyl substituted with 0-2 R^{10a} and piperidinyl
 substituted with 0-2 R^{10a};
 each R^{10a} is independently a member selected from the group consisting of Cl, F, C₁-
 C₄ alkyl, C₁-C₄ alkoxy, OCF₃, and CF₃;
 alternatively, two R^{10a} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical comprises 1 or 2 heteroatom(s);
 Z is phenylene;
 R¹¹ is a member selected from the group consisting of indolyl substituted with 0-2
 R^{11a}, benzofuranyl substituted with 0-2 R^{11a}, benzothieryl substituted with 0-2

R^{11a} , benzothiazole substituted with 0-2 R^{11a} , benzisoxazolyl substituted with
 0-2 R^{11a} , benzoxazolyl substituted with 0-2 R^{11a} , and pyrazolo[1,5-
 a]pyrimidinyl substituted with 0-2 R^{11a} , piperidinyl N-substituted with 0-1
 R^{11b} , morpholinyl N-substituted with 0-1 R^{11b} ; and 2-oxo-pyrrolidinyl with 0-1
 R^{11b} ;
 each R^{11a} is independently a member selected from the group consisting of Cl, F,
 NH_2 , CH_3 , OCH_3 , $-C(=O)OCH_3$, OCF_3 , and CF_3 ;
 each R^{11b} is independently a member selected from the group consisting of
 pyrimidinyl substituted with 0-2 R^{11c} ; benzyl, acetyl, CH_2 -furanyl, and CH_2 -
 thienyl;
 each R^{11c} is independently a member selected from the group consisting of Br and
 CH_3 ;
 R^{12} is $(CH_2)_s R^{12a}$;
 R^{12a} is a member selected from the group consisting of cyclopentyl and cyclohexyl;
 s is the integer 1 or 2;
 R^{13} is a member selected from the group consisting of H and C_1 - C_4 alkyl;
 R^{14} is pyrimidinyl substituted with 0-2 R^{16} ;
 R^{15} is a member selected from the group consisting of C_1 - C_4 alkyl, morpholinyl,
 pyrrolidinyl and piperidinyl;
 R^{16} is a member selected from the group consisting of CH_3 and OCH_3 ;
 each of R^{17} and R^{18} is independently a member of H, OH, F, phenyl and C_1 - C_3 alkyl;
 alternatively, R^{17} and R^{18} may be taken together to form a C_3 - C_6 cycloalkyl;
 Ar is a phenyl substituted with 0-2 R^{19} ;
 each R^{19} is independently a member selected from the group consisting of F, Cl,
 $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 ;
 and a pharmaceutically acceptable excipient.

10. The composition of claim 9, wherein said compound is a member
 selected from the compounds of Table I.

11. A method of selectively inhibiting cathepsin S activity in a mammal in
 need thereof, comprising administering to said mammal a therapeutically effective amount of
 a compound of Formula I:



or a pharmaceutically acceptable salt or prodrug thereof,

wherein:

W is a member selected from the group consisting of

$\text{R}^1\text{-X-(C=O)-NH-CHR}^2\text{-}$,

$\text{R}^4\text{-Y-(C=O)-NH-CHR}^3\text{-}$,

$\text{R}^6\text{-(C=O)-NH-CHR}^5\text{-}$,

$\text{R}^7\text{-NH-(C=O)-NH-CHR}^8\text{-}$,

$\text{R}^{10}\text{-Z-(C=O)-NH-CHR}^9\text{-}$, and

$\text{R}^{11}\text{-(C=O)-NH-CHR}^{12}\text{-}$;

R^1 is a member selected from the group consisting of phenyl substituted with 0-2 R^{1a} ,
pyridyl substituted with 0-2 R^{1a} , and pyridinium N-oxide substituted with 0-2
 R^{1a} ;

each R^{1a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

X is a member selected from the group consisting of furanylene substituted with 0-1
 R^x , thienylene substituted with 0-1 R^x , pyrazolyene substituted with 0-1 R^x ,
thiazolyene substituted with 0-1 R^x , and oxazolyene substituted with 0-1 R^x ;

R^x is a member selected from the group consisting of F, Cl, CH_3 and CF_3 ;

R^2 is a member selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
and $(\text{CH}_2)_n\text{R}^{2b}$;

each R^{2a} is independently a member selected from the group consisting of Cl, F,
 OCF_3 , OCH_3 , CH_3 and CF_3 ;

R^{2b} is independently a member selected from the group consisting of phenyl
substituted with 0-2 R^{2a} ; cyclopentyl, cyclohexyl and tetrahydropyranyl;

n is the integer 1 or 2;

R^3 is $(\text{CH}_2)_m\text{R}^{3b}$;

R^{3b} is selected from the group consisting of phenyl substituted with 0-2 R^{2a} ,
cyclopentyl and cyclohexyl;

m is the integer 1 or 2;

R^4 is a member selected from the group consisting of phenyl substituted with 0-3 R^{4a} ,
thienyl, tetrazolyl, cyclopentenyl and indolyl;

each R^{4a} is a member selected from the group consisting of phenyl, OH, C_1 - C_4 alkyl,
 C_1 - C_4 alkoxy, CF_3 , OCF_3 , F, Cl, $CH_3S(=O)_2$ -, morpholinyl, pyrrolidinyl,
 piperidinyl and 4-acetypiperazinyl;
 Y is a member selected from the group consisting of $-CR^{17}R^{18}$, $-NH-CH_2-$ and $-O-$
 CH_2- ;
 R^5 is a member selected from the group consisting of phenyl substituted with 0-2 R^{5a} ,
 thiophene, naphthyl, and CH_2R^{5b} , CH_2CH_2 (cyclohexyl),
 $CH_2CH_2CH_2$ (cyclohexyl), CH_2CH_2Ph , $CH(CH_3)R^{5c}$, $CH_2CH=CHPh$, -
 CH_2OCH_2Ph , $-CH(CH_3)OCH_2Ph$;
 each R^{5a} is independently a member selected from the group consisting of F, Cl, NO_2 ,
 OCH_3 , OCH_2Ph , OPh , CH_3 , OCF_3 and CF_3 ;
 R^{5b} is independently a member selected from the group consisting of phenyl
 substituted with 0-2 R^{5c} ; cyclopentyl, cyclohexyl, naphthyl, indolyl and
 pyridyl;
 R^{5c} is independently a member selected from the group consisting of OH, Cl, F, Br, I,
 CN, NO_2 , CH_3 , OCH_3 , tBu , $O-tBu$, $-NHC(=O)CH_3$, CF_3 , OCF_3 ; phenyl
 substituted with 0-2 R^{5d} ; phenoxy substituted with 0-2 R^{5d} ; benzyloxy
 substituted with 0-2 R^{5d} ; pyridyl substituted with 0-2 R^{5d} ; pyrimidinyl
 substituted with 0-2 R^{5d} ; thienyl substituted with 0-2 R^{5d} ;
 R^{5d} is independently a member selected from the group consisting of CH_3 , Cl, F,
 OCH_3 , CF_3 , OCF_3 , $N(CH_3)_2$, acetyl, OH, CH_2OH , NH_2 , CN and NO_2 ;
 R^{5e} is phenyl substituted with 0-2 R^{5a} ;
 R^6 is a member selected from the group consisting of phenyl substituted with 0-3 R^{6a} ,
 furanyl substituted with 0-2 R^{6b} , thienyl substituted with 0-2 R^{6b} , oxazolyl
 substituted with 0-2 R^{6b} , thiazolyl substituted with 0-2 R^{6b} , pyridyl,
 pyridazinyl and cyclopropyl;
 each R^{6a} is independently a member selected from the group consisting of Cl, F, Br,
 OCF_3 , CF_3 , C_1 - C_4 alkyl, C_1 - C_4 alkoxy, $-S(=O)_2CH_3$, CN, $-N(CH_3)_2$, OCF_2H , -
 CH_2 -benzimidazole, $-NH-S(=O)_2CH_3$, $-NR^{13}R^{14}$, OR^{14} , CH_2 -morpholine,
 CH_2NH_2 , OCH_2Ph , and OPh ;
 alternatively, two R^{6a} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical has 1 or 2 oxygen atom(s);

each R^{6b} is independently a member selected from the group consisting of NH_2 , F, Cl,
 Br, $-S(=O)_2R^{15}$, CH_3 , and CF_3
 R^7 is a member selected from the group consisting of $(CH_2)_p R^{7a}$, and naphthyl
 substituted with 0-2 R^{7b} ;
 p is the integer 0, 1, or 2;
 R^{7a} is phenyl substituted with 0-2 R^{7b} ;
 R^{7b} is a member selected from the group consisting of F, Cl, CF_3 , C_1 - C_4 alkyl, C_1 - C_4
 alkoxy, OCF_3 , phenoxy and acetyl;
 alternatively, two R^{7b} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical has 1 or 2 oxygen atom(s);
 R^8 is $-CH_2-R^{3b}$;
 R^9 is $(CH_2)_q R^{9a}$;
 R^{9a} is a member selected from the group consisting of cyclopentyl, phenyl and
 cyclohexyl;
 q is the integer 1 or 2;
 R^{10} is a member selected from the group consisting of phenyl substituted with 0-2
 R^{10a} , 5 membered heteroaryl containing 1 to 4 heteroatoms each independently
 a member selected from the group consisting of N, O and S, wherein said
 heteroaryl is substituted with 0-2 R^{10a} , 6 membered heteroaryl containing 1 to
 2 N, wherein said heteroaryl is substituted with 0-2 R^{10a} , morpholinyl
 substituted with 0-2 R^{10a} , piperazinyl substituted with 0-2 R^{10a} and piperidinyl
 substituted with 0-2 R^{10a} ;
 each R^{10a} is independently a member selected from the group consisting of Cl, F, C_1 -
 C_4 alkyl, C_1 - C_4 alkoxy, OCF_3 , and CF_3 ;
 alternatively, two R^{10a} substituents on adjacent atoms may be combined to form a 5 to
 6 membered heterocyclic fused radical, wherein said 5 to 6 membered
 heterocyclic fused radical comprises 1 or 2 heteroatom(s);
 Z is phenylene;
 R^{11} is a member selected from the group consisting of indolyl substituted with 0-2
 R^{11a} , benzofuranyl substituted with 0-2 R^{11a} , benzothienyl substituted with 0-2
 R^{11a} , benzothiazole substituted with 0-2 R^{11a} , benzisoxazolyl substituted with
 0-2 R^{11a} , benzoxazolyl substituted with 0-2 R^{11a} , and pyrazolo[1,5-
 a]pyrimidinyl substituted with 0-2 R^{11a} , piperidinyl N-substituted with 0-1

103 R^{11b} , morpholinyl N-substituted with 0-1 R^{11b} ; and 2-oxo-pyrrolidinyl with 0-1
104 R^{11b} ;
105 each R^{11a} is independently a member selected from the group consisting of Cl, F,
106 NH_2 , CH_3 , OCH_3 , $-C(=O)OCH_3$, OCF_3 , and CF_3 ;
107 each R^{11b} is independently a member selected from the group consisting of
108 pyrimidinyl substituted with 0-2 R^{11c} ; benzyl, acetyl, CH_2 -furanlyl, and CH_2 -
109 thienyl;
110 each R^{11c} is independently a member selected from the group consisting of Br and
111 CH_3 ;
112 R^{12} is $(CH_2)_s R^{12a}$;
113 R^{12a} is a member selected from the group consisting of cyclopentyl and cyclohexyl;
114 s is the integer 1 or 2;
115 R^{13} is a member selected from the group consisting of H and C_1 - C_4 alkyl;
116 R^{14} is pyrimidinyl substituted with 0-2 R^{16} ;
117 R^{15} is a member selected from the group consisting of C_1 - C_4 alkyl, morpholinyl,
118 pyrrolidinyl and piperidinyl;
119 R^{16} is a member selected from the group consisting of CH_3 and OCH_3 ;
120 each of R^{17} and R^{18} is independently a member of H, OH, F, phenyl and C_1 - C_3 alkyl;
121 alternatively, R^{17} and R^{18} may be taken together to form a C_3 - C_6 cycloalkyl;
122 Ar is a phenyl substituted with 0-2 R^{19} ; and
123 each R^{19} is independently a member selected from the group consisting of F, Cl,
124 $COOH$, C_1 - C_4 alkoxy, $OCHF_2$ and OCF_3 .

1 12. The method of claim 11, wherein the cathepsin S inhibition constant
2 for a compound of Formula I is less than 10 μM .

1 13. The method of claim 12, wherein the cathepsin S inhibition constant
2 for a compound of Formula I is less than 1.0 μM .

1 14. The method of claim 13, wherein the cathepsin S inhibition constant
2 for a compound of Formula I is less than 0.1 μM .

1 15. The method of claim 11, wherein cathepsin S is selectively inhibited in
2 the presence of cathepsin K.

1 **16.** The method of claim **15**, wherein the inhibition constant of a
2 compound of Formula I for cathepsin K is at least 10 times greater than a cathepsin S
3 inhibition constant of a compound of Formula I.

1 **17.** The method of claim **16**, wherein the inhibition constant of a
2 compound of Formula I for cathepsin K is at least 100 times greater than said cathepsin S
3 inhibition constant of a compound of Formula I.

1 **18.** The method of claim **17**, wherein the inhibition constant of a
2 compound of Formula I for cathepsin K is at least 1000 times greater than said cathepsin S
3 inhibition constant of a compound of Formula I.

1 **19.** The method of claim **11**, wherein said compound is a member selected
2 from the compounds of Table I.